



# ***Emerging Contaminants and the Ramifications for Environmental Science***

**Christian G. Daughton, Ph.D.**

*Chief, Environmental Chemistry Branch*

Environmental Sciences Division

National Exposure Research Laboratory

Office of Research and Development

Environmental Protection Agency

Las Vegas, Nevada 89119

[daughton.christian@epa.gov](mailto:daughton.christian@epa.gov)

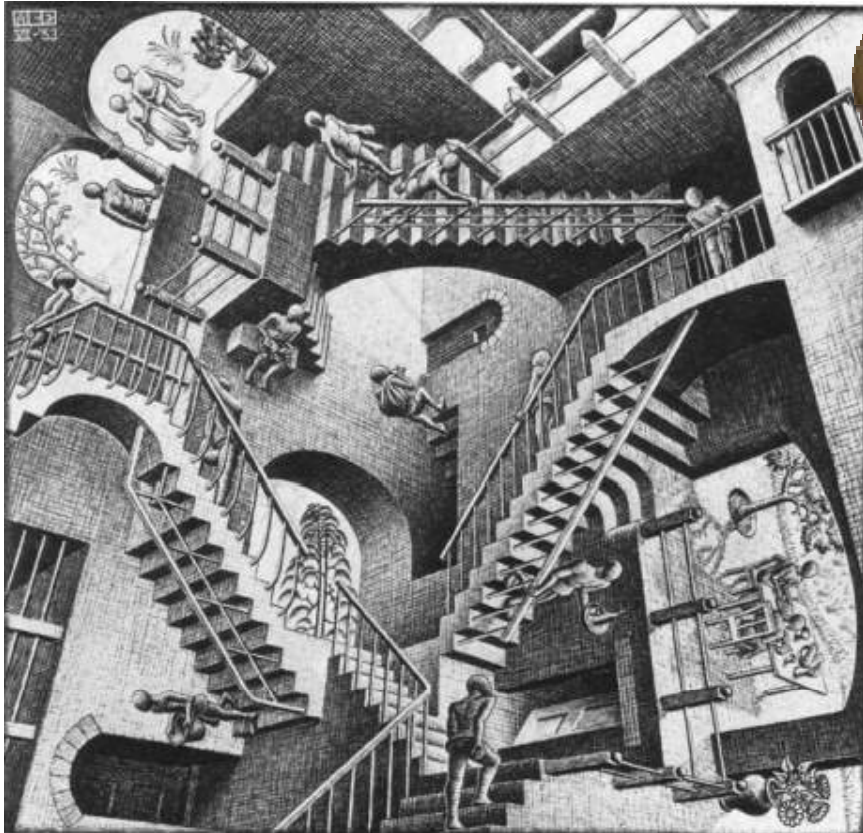
**Office of Research and Development**

National Exposure Research Laboratory, Environmental Sciences Division, Las Vegas, Nevada

**Does anyone know what an  
*Emerging Contaminant* really is?**



# ***Emerging Contaminants ...***



***... depend on  
one's perspective***

# What Makes a “Contaminant of Emerging Concern”?

*Emerging contaminants* (ECs) include more than simply chemicals previously not known to occur in the environment.

ECs also include chemicals already known to occur but now displaying new characteristics not previously suspected or recognized, such as those involving:

- origin or source (e.g., via sweat and dermal transfer)
- location (e.g., “out-of-place” chemicals; “chemical weeds”)
- unusual concentrations or levels  
(e.g., enriched by sorption to plastics in oceans)
- transformation and fate pathways
- exposure routes
- biological effects pathways or endpoints



## *Slicing the Pollutant Pie*

Environmental contaminants can be categorized across many dimensions.



A host of acronyms describe different slices, often breeding confusion.

# *Acronym Anarchy*

EDCs – CMRs – PBTs – vPvB – POPs  
– OWCs – PPCPs – PhACs – EPOCs  
– COCs – COPCs – CECs – XOCs –  
HPVs – PDPs – HDPs – POHOs –  
SVHCs

ppm – ppb – ppt – ppq  
micro – nano – pico

*... leads to acronym stupefaction*



## The Confusing Terminology of Chemical Pollutant Classes

<b>Class or Grouping</b>	<b>Grouped According to:</b>
<b>EDC</b> (Endocrine Disrupting Chemical) <b>CMR</b> (Carcinogenic, Mutagenic, toxic to Reproduction)	toxicological mode of action or endpoint
<b>PBT</b> (Persistent, Bioaccumulative Toxic) <b>vPvB</b> (very Persistent, very Bioaccumulative) <b>POP</b> (Persistent Organic Pollutant)	environmental properties
<b>micro-pollutants; micro-constituents</b> <b>trace pollutants/residues</b>	frequency/level of occurrence
<b>OWCs</b> (Organic Wastewater Contaminants)	location of occurrence
<b>PPCPs</b> (also PhACs, which is a subset)	type of intended usage
<b>priority pollutants</b> and others	regulation
<b>unregulated pollutants</b>	absence of regulation
<b>“chemical weeds”</b>	public perception ("out-of-place chemicals")
<b>emerging contaminants/pollutants</b> <b>EPOCs</b> (emerging pollutants of concern) <b>COCs</b> (chemicals of concern) <b>COPCs</b> (chemicals of potential concern) <b>CECs</b> (chemicals of emerging concern)	novelty, fad, timeliness, or new concern
<b>xenobiotics, exotics</b> <b>xenobiotic organic compounds (XOCs)</b>	foreign versus endogenous
<b>toxicants, toxins, toxics, perturbogens</b> (agonists, antagonists, activators, repressors, inhibitors, regulators, modulators)	overall toxicity ( <i>note</i> : "toxins" comprise a special subset of toxicants - those naturally synthesized, primarily proteins; "toxics" is jargon for "toxicants")
<b>HPV</b> (high production volume) chemicals	quantity (manufactured/imported in US $\geq 1$ million pounds/year)
<b>PDPs</b> or <b>HDPs</b> (population- or human-derived pollutants/constituents) <b>POHO</b> (pollutants of human origin)	source or origin

Adapted and extended by Daughton from: *Renewable Resources Journal*, 2005, 23(4):6-23.

# Context of “*Emerging*”

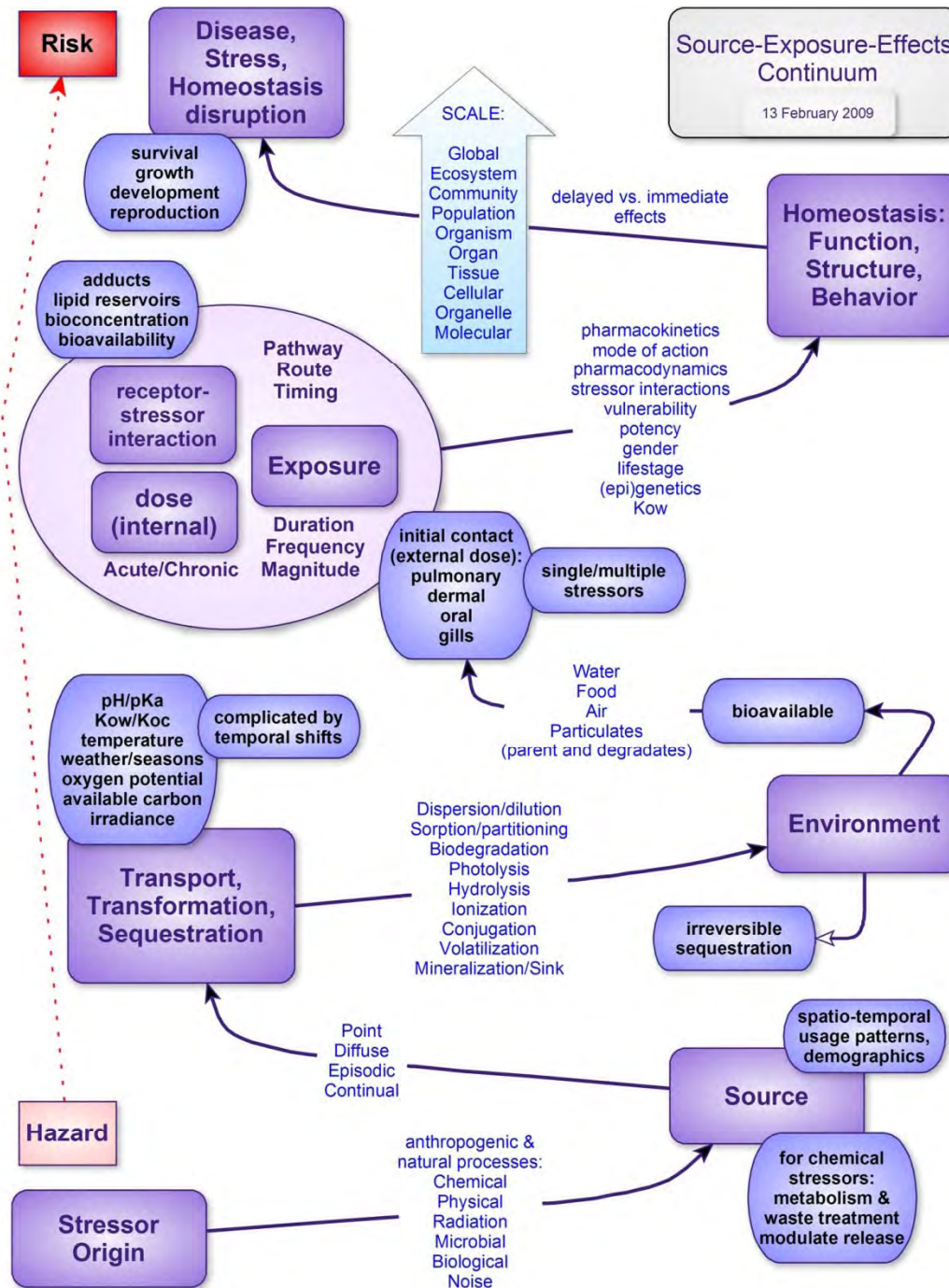
The term "emerging" can be applied to any part of the continuum of the risk paradigm – spanning stressors to exposure and effects. It applies not just to chemicals, but also to any previously unrecognized, unexpected, or under-appreciated aspects of environmental science.

The study of "emerging" contaminants is really one of more carefully examining the interface between humans and the environment; and of continually expanding the conventional boundaries surrounding all facets of the source-to-effects continuum.

The term "ECs" causes confusion because it pertains not just to chemicals themselves but also to the many dimensions associated with their properties and the source-to-effects continuum.







## *Some EPA Perspective*

Statement of **Peter S. Silva** (confirmed, June 2009, as Assistant Administrator for the Office of Water) at hearing of Nominations Committee on Environment and Public Works United States Senate:

"... despite the considerable progress we have made in the last three decades, we now see additional challenges have arisen in the areas of non-point source pollution and in new emerging pollutants of concern." (12 May 2009)

"Managing Chemical Risks" is one of EPA's five new priority areas identified by EPA's new Administrator, **Lisa Jackson**:

"It is clear that we are not doing an adequate job of assessing and managing the risks of chemicals in consumer products, the workplace and the environment." (23 January 2009)

## *Some EPA Perspective*

**Peter S. Silva** (American Water Works Association, Annual Conference and Exposition, 15 June 2009, San Diego)

"There is a growing nexus between the way we supply water and how we face environmental impacts."

"Even direct potable reuse is being considered."



*ECs will play prominent roles in these efforts.*

# "Emerging" Contaminants

**Most "emerging" contaminants are not new to the environment**



- **Two major sources for contaminants that are truly "new" to the environment:**
  - **Chemicals newly introduced to commerce** (e.g., new approved or unapproved drugs, pesticides, nanomaterials).
  - **By-products of new anthropogenic processes** (e.g., nanotechnology).
- **Previously unrecognized pollutants can come to our attention as a result of:**
  - **New advances in chemical analysis** (e.g., "non-target" identification).
  - **Ability to detect existing pollutants at ever-lower concentrations** (single-molecule detection is now possible).
  - **Exploring exposure sources not previously examined** (e.g., certain foods as a significant source of acrylamide; estrogens in plastics; PPCPs in biosolids; alkaloids in pyrolysis products from controlled forest burns).

# examples

Old pollutants in new places: organohalogens in dolphin & seal brain; conventional POPs and their metabolites in tissues never before explored.

Old chemicals as new suspects: tungsten: mono-, poly- & heteropoly-tungstates

Toxicants masquerading as "inert" ingredients: polyethoxylated tallow amine used as adjuvant for formulating glyphosate; shows high aquatic toxicity and toxicity in human cell cultures.

Occurring at levels or locations never before observed or suspected: APIs in drug manufacturing waste streams. PPCPs in household graywater. Lipophilic contaminants sorbed to plastics in oceans.

New pollutants: perfluorinated phosphonic acids (in waters); unapproved drugs

Source or origin: marine production of halogenated organics; alkaloids in particulates/aerosols from controlled burns in forests (pyrolytic conditions); melamine in infant formula, originating as metabolite of cyromazine, an insecticide used on crops and forage.

Route of transport to the environment (contributing to exposure risk): excretion of drugs via the skin (sweat); illicit drugs in ambient air.

Exposure pathway: interdermal transfer among humans.



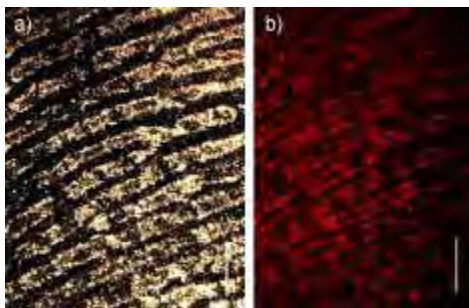


# *Emerging Contaminants: Humanity's Chemical Footprints and Fingerprints*

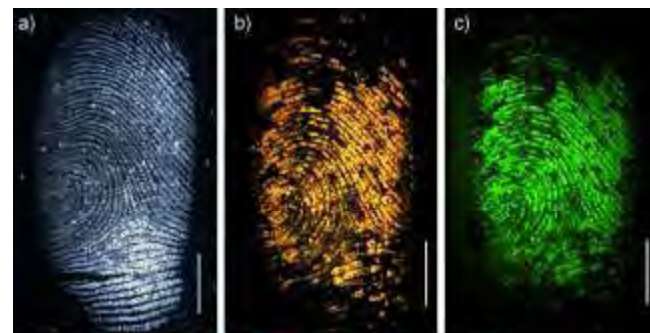
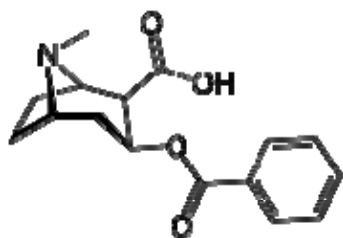
- Enter the environment by way of diffuse, complex, and interconnected networks fed by myriad numbers of people.
- ECs reflect:
  - The intimate, inseparable, and immediate connections of humans and the environment.
  - The combined actions, behaviors, and activities of individuals.
  - The extensive interconnections and feedback loops between humans and the environment.
- ECs serve as direct measures of the types, extent, and magnitude of these connections.



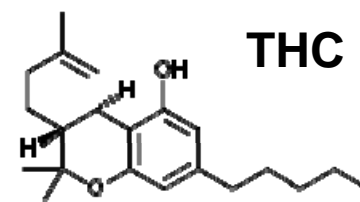
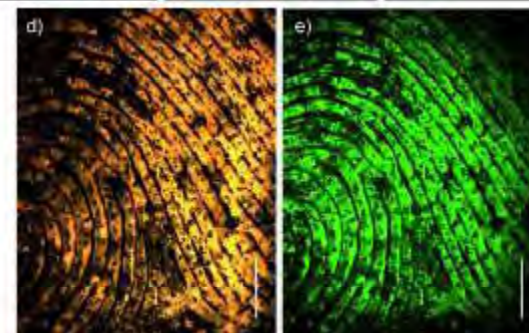
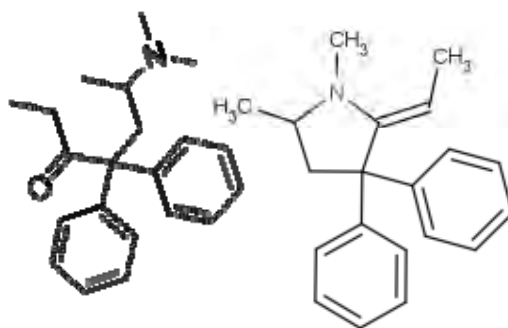
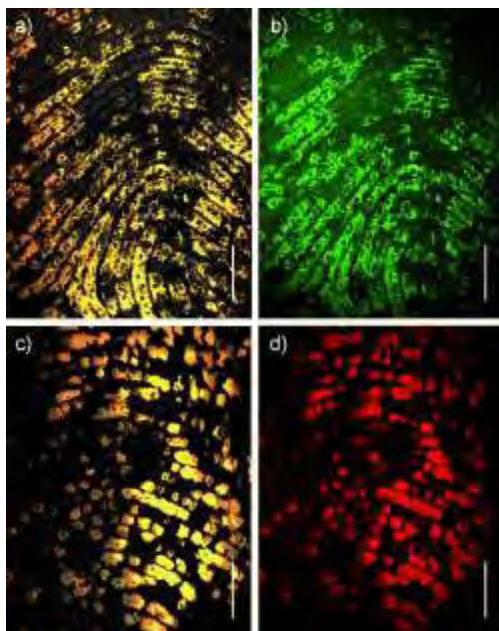
# Fingerprints as contributors of environmental contaminants



**benzoylecgonine**



**methadone & EDDP**



**THC**

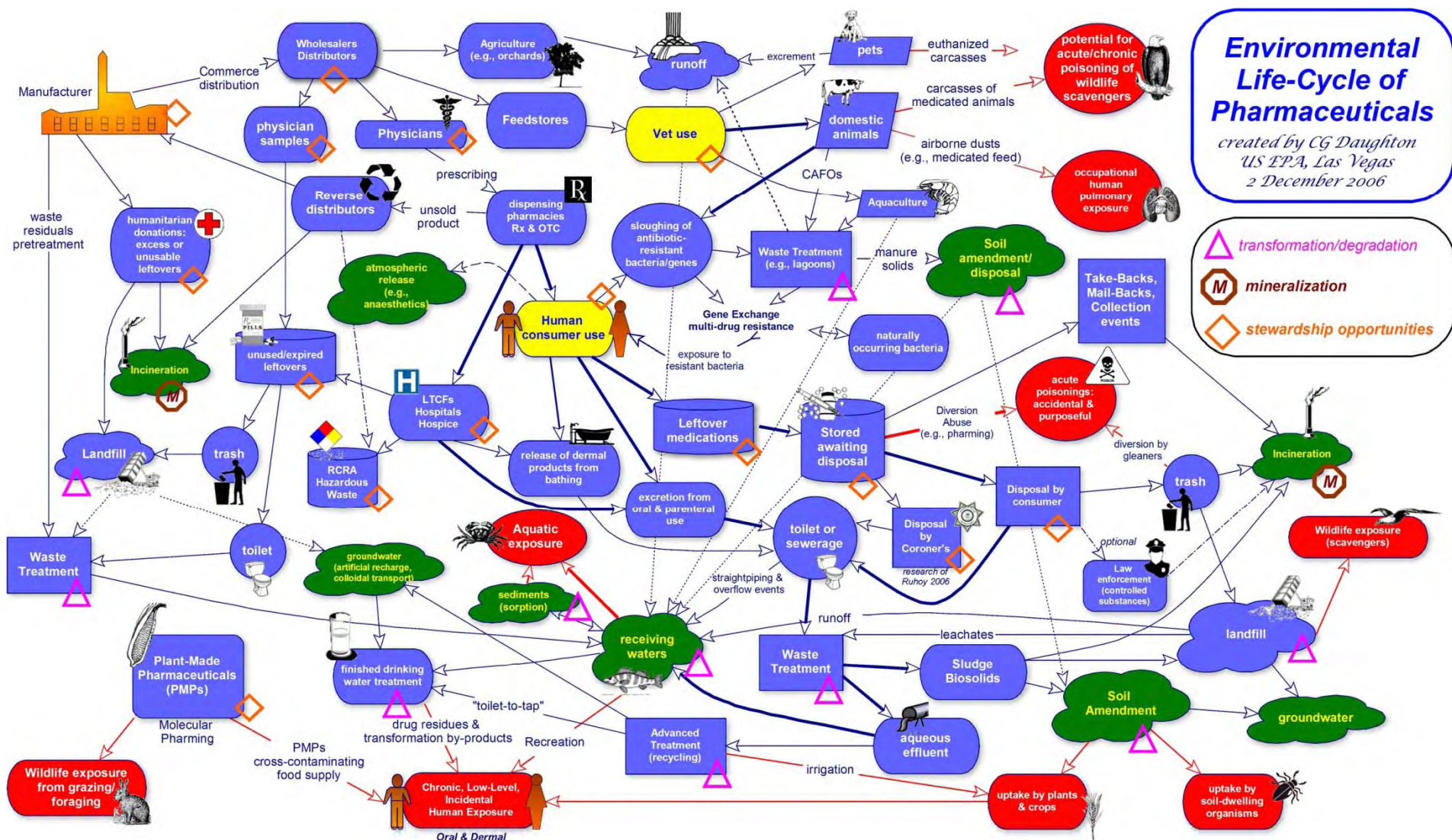
*benzoylecgonine* (major cocaine metabolite)

*EDDP*: 2-ethylidene-1,5-dimethyl-3,3-diphenylpyrrolidine (major methadone metabolite)

*THC*:  $\Delta^9$ -tetrahydrocannabinol (main psychoactive of marijuana)

Hazarika P, Jickells SM, Wolff K, Russell DA (2008) Imaging of Latent Fingerprints through the Detection of Drugs and Metabolites. *Angew Chem Int Ed* 47:10167-10170.





*Challenges for environmental chemical analysis:  
Complexities in identifying and monitoring chemical unknowns*



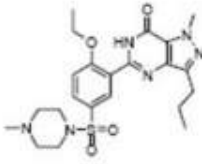
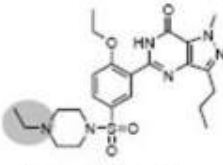
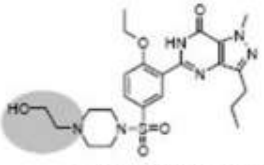
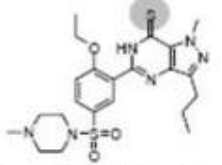
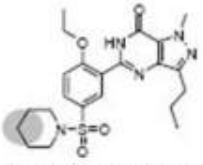
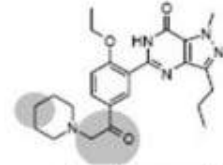
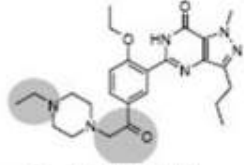
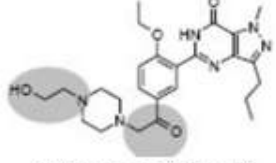
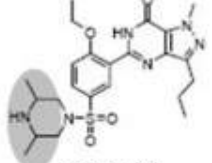
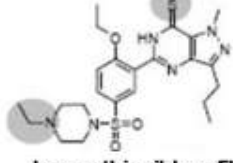
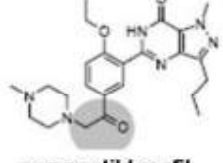
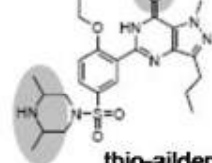
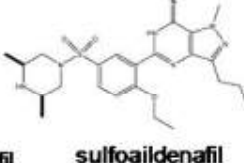
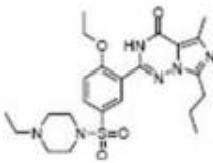
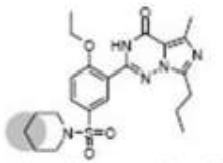
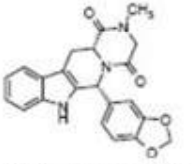
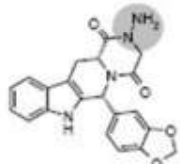
- **Chiral enantiomers**: long proven important for certain pharmaceuticals and pesticides (e.g., improved profiles for effectiveness and toxicity versus hydrogen analogs). Subject to enantiomeric enrichment via microbial degradation.
- **Nanomaterials**: pose enormous challenges for chemical characterization because of diversity of structures and possible complexity of weathered products.
- **Unapproved drugs**: unapproved analogs of registered drugs; sold OTC in adulterated herbal supplements and in fraudulent prescription medications.
- **Mechanical bonds**: rotaxanes, catenanes, molecular knots

# Designer Drugs: Environmental Contaminant Unknowns

Undeclared drug active ingredients (new molecules) – known or unknown to FDA.  
Adulterants in herbal supplements or OTC/Rx drugs, often at high levels.

*Example:* analogs of the approved phosphodiesterase type 5 (PDE-5) inhibitors (used primarily in the treatment of erectile dysfunction), such as sildenafil, vardenafil, and tadalafil.

B.J. Venhuis, D.M. Barends, M.E. Zwaagstra, and D. de Kaste "Recent developments in counterfeits and imitations of Viagra, Cialis and Levitra: A 2005-2006 update," 2007, RIVM (Netherlands National Institute for Public Health and the Environment), RIVM Report 370030001/2007, Bilthoven, the Netherlands, 61 pp;  
<http://www.rivm.nl/bibliotheek/rapporten/370030001.pdf>

Registered APIs	Analogues identified		
 sildenafil (Viagra®)	 homosildenafil	 hydroxyhomosildenafil	 thiosildenafil
	 piperidino-sildenafil	 piperidino-acetildenafil	 acetildenafil
	 hydroxy-acetildenafil	 aildenafil	 homo-thiosildenafil
	 nor-acetildenafil	 thio-aildenafil	 sulfoaildenafil
 vardenafil (Levitra®)	 piperidino-vardenafil		
 tadalafil (Cialis®)	Herbal supplements: Zencore Tabs LibieXtreme Libimax X  amino-tadalafil		

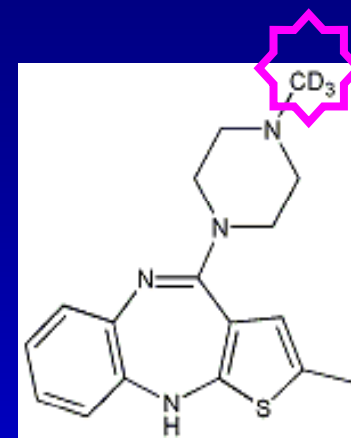
Herbal supplements:  
Y-4ever  
Powermania



## Challenges for environmental chemical analysis: Complexities in identifying and monitoring chemical unknowns

### ■ **Isotopic substitution:** *deuterated analogs* of pharmaceuticals and pesticides long known for altered pharmacokinetics/pharmacodynamics because of kinetic isotope effect.

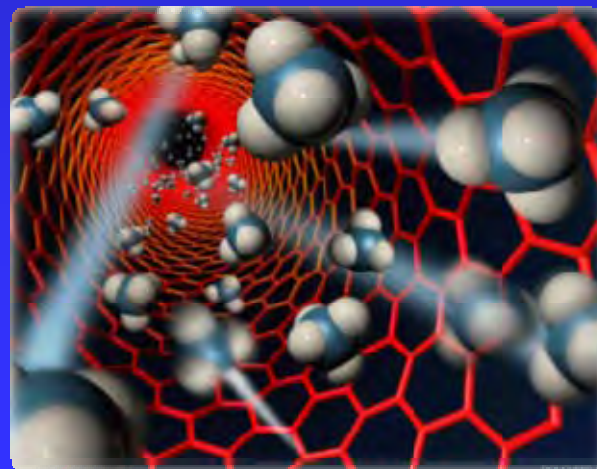
- Pharmacologic deuteration yields drugs with greater stability. Can facilitate longer half-lives (increased duration of action from hindered first-pass metabolism); fewer side effects from lower dose, "metabolic switching," and reduced drug-drug interactions.
- Major unknown for deuterated drugs: alteration of environmental fate & transport?
- Major new analytical challenge: each API can have one or more deuterated analogs, increasing the numbers of potential analytes. But moreover, these analogs would greatly hinder the identification of each other if they cannot be physically separated prior to mass spectral detection (a result of radically altered isotopic abundances).



# ***Structurally Undefinable Xenobiotics***

## **Importance of Developing the Means to Evaluate and Monitor "Undefinable" Xenobiotics**

- The widespread emergence in consumer products of new **nanomaterials** poses a host of currently unanswerable questions regarding their environmental chemistry and environmental toxicology.
- In particular, the natural weathering or environmental transformation of these materials holds the potential to yield numerous additional, **structurally undefinable products**.
- **Nanomaterials pose numerous, currently intractable challenges to their monitoring in the environment.**



# It's Been Only 10 Years

The first published documents to prominently feature the term *emerging contaminants/pollutants* were probably the 1999 report from the 1998 NRC workshop ("Identifying Future Drinking Water Contaminants") and several 1999 government reports by USGS.

The first publication in a journal was in 2001:  
Daughton, C.G. "Emerging Pollutants, and Communicating the Science of Environmental Chemistry and Mass Spectrometry: Pharmaceuticals in the Environment," *J. Am. Soc. Mass Spectrom.* **2001**, 12(10), 1067-1076; doi: 10.1016/S1044-0305(01)00287-2; available:  
[http://www.epa.gov/nerlesd1/bios/daughton/book\\_jasma.pdf](http://www.epa.gov/nerlesd1/bios/daughton/book_jasma.pdf)



# *Emerging Contaminants*

## - Status Check -

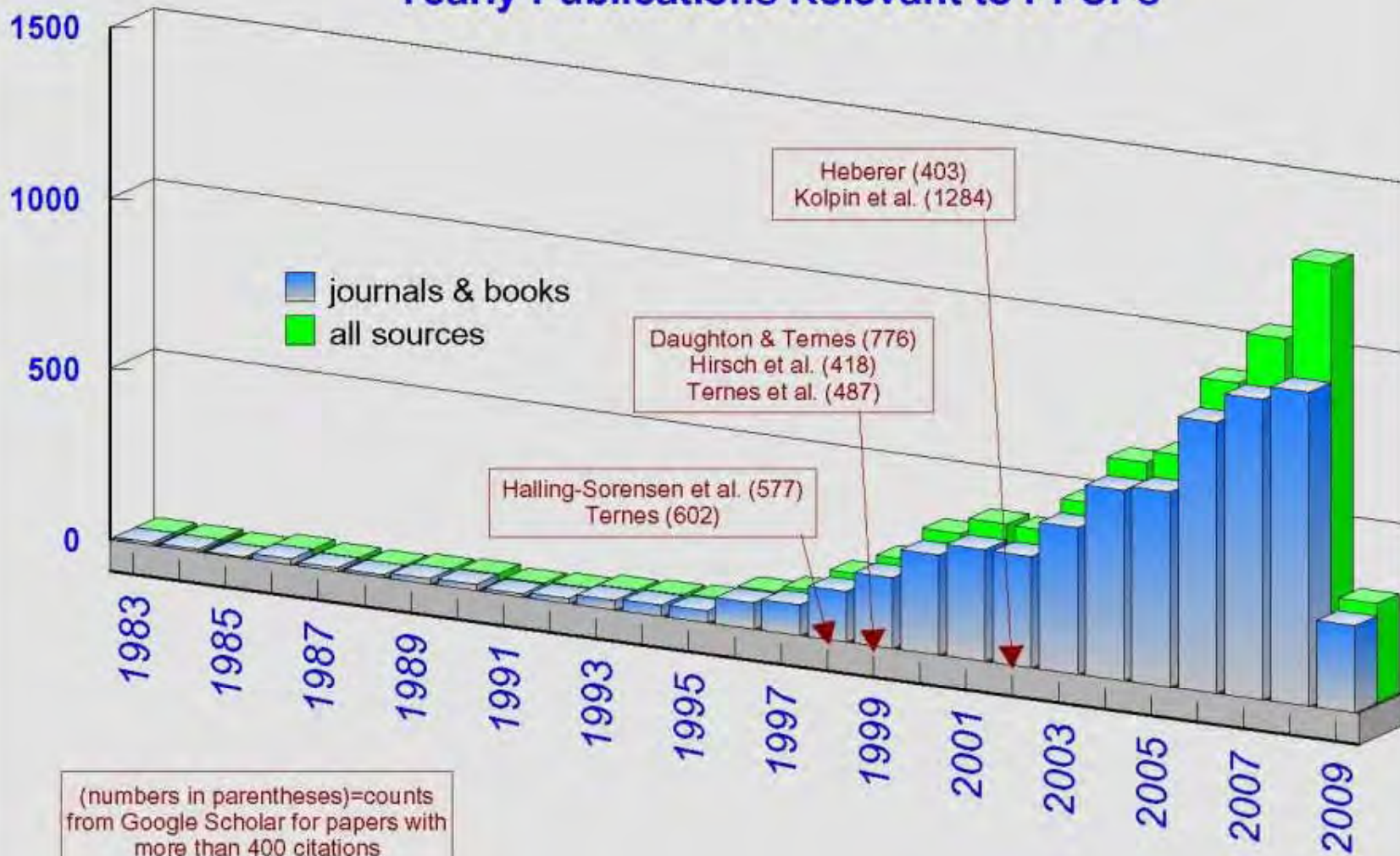


Passage of a decade marks an excellent time to reflect on a number of questions surrounding this rather ill-defined but broadly used term.

- Has the term served us well?
- Do we share a common understanding as to what an "emerging" contaminant might be?
- Is it leading us where we need to be in terms of a scientific discipline?
- What message does it convey to the public, legislators, and regulators?
- Are we perhaps suffering from its overuse – having become an obligatory adjective?
- What about the galaxies of other chemicals that currently evade detection and cannot yet be classified as “emerging”?



## Yearly Publications Relevant to PPCPs



note: data for 2009 only through first 8 weeks

# A More Accurate Birth Date – 150 years ago

George P. Marsh: forerunner to the modern environmental movement. Over 150 years ago, recognized that interconnections pervade all of nature.

Birth of the idea of incremental, cumulative impacts from small forces. Foretold the new paradigm of countless point-sources combining to serve as a unified dispersed source: **the significance of the combined but miniscule actions, activities, and behaviors of multitudes of individuals.**

Perhaps the first scientific examination and documentation of the thesis that harm accumulates, regardless of the seeming insignificance of each act of harm.



Marsh GP. 1864. *Man and nature; or, Physical geography as modified by human action.* C. Scribner, New York.

## 1864: Birth of the idea of incremental, cumulative impacts [or ... *Small Can Be as Big as Big*]

From the chapter:

**"Nothing Small in Nature."**

Marsh GP. 1864.

Man and nature; or, Physical geography as modified by human action.

C. Scribner, New York.



“No atom can be disturbed in place, or undergo any change of temperature, of electrical state, or other material condition, without affecting, by attraction or repulsion or other communication, the surrounding atoms. These, again, by the same law, transmit the influence to other atoms, and the impulse thus given extends through the whole material universe.” “...reaction does not restore disturbed atoms to their former place and condition, and consequently the effects of the least material change are never cancelled, but in some way perpetuated, so that *no action can take place in ... nature, without leaving all matter in a different state from what it would have been if such action had not occurred.*”

## *Emerging Challenges from Emerging Contaminants*

- Increasingly advanced methods of analysis allow peering into the shadows of chemical space with ever-greater magnification and clarity.
- At ever-lower exposure levels, it may no longer be possible to deconvolute a purported effect from its incidence as ambient (natural) background.
- Ever-lower detection limits pose increasingly greater challenges for assessing, communicating, and ameliorating ever-diminishing risks.
- Where do these risks reside within the overall scope of larger environmental concerns?



# Complex Mixtures of Chemical Stressors at Ultra-Low Concentrations:

*Pose Major Challenges for Measurement and for  
Assessing Exposure and Biological Effects*



***What portion of overall risk is  
contributed by emerging  
contaminants?***

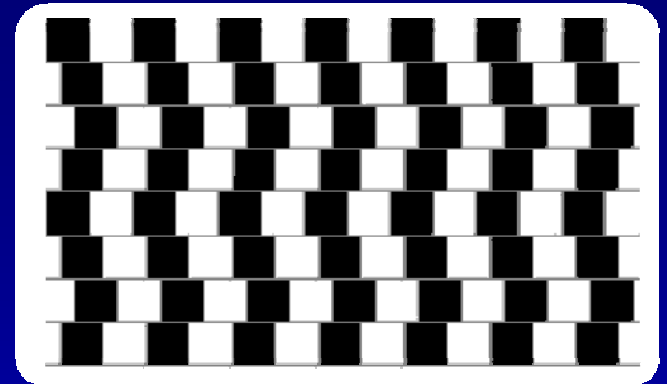




## *Public Perception*

Public perceives only two reasons that ECs occur in the environment:

1. Newly introduced to commerce
  2. We failed to detect them up until now
- ECs serve to highlight the fact that the approach to environmental monitoring is reactive rather than anticipatory.
  - The term “emerging” can alarm the public, possibly cultivating a negative impression of failure on the part of environmental scientists (but perception does not align with reality).
    - After all, the sole reason that a pollutant becomes known as "emerging" in the first place is simply because of our failure to anticipate or predict (in an ideal world, we should be able to identify and prevent the "emergence" of chemicals).
  - Problem is exacerbated with ongoing identification of yet more ECs - with few actions to mitigate or prevent future ones.





# The Chemical Universe

We live in a chemical sea of continually changing composition – comprising both anthropogenic and naturally occurring chemical stressors.

Unlike biota, chemical pollutants have no boundaries in their global distribution –  
“*everything is everywhere,*” only the concentrations vary.



# The Chemical Universe

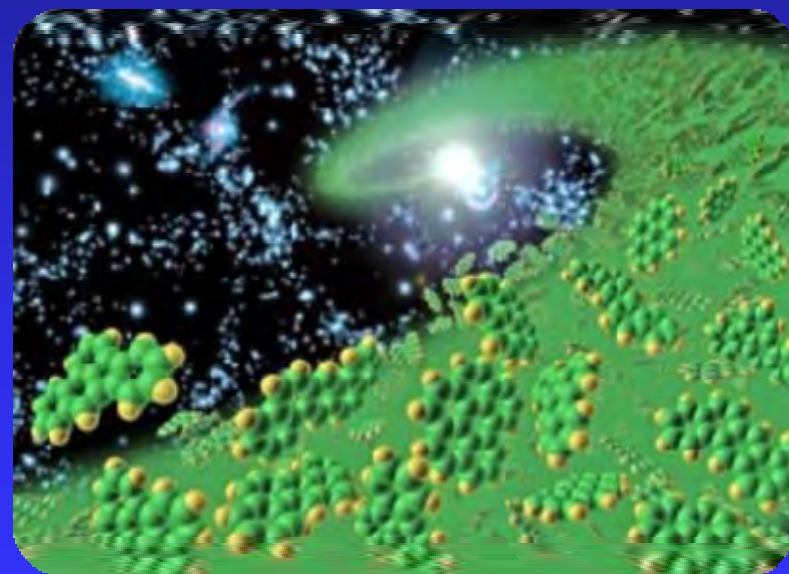
## The *KNOWN* Universe

As of April 2010, over 52 million organic and inorganic substances had been assigned CAS RNs.

(indexed by the American Chemical Society's Chemical Abstracts Service in their CAS Registry; excluding bio-sequences such as proteins and nucleotides:

<http://www.cas.org/expertise/cascontent/registry/regsys.html>)

- Of these millions of known chemicals, nearly 40 million were commercially available.
- Roughly only 280,000 are inventoried or regulated by government bodies worldwide - - representing only 0.7% of those that are commercially available or less than 0.6% of the known universe of chemicals.
- Approximately 12,000 new substances are added each day.



# The Chemical Universe

## The *POTENTIAL* Universe

While the *KNOWN* universe of commercial chemicals might seem large (52 million), the universe of *POTENTIAL* chemicals (those that could possibly be synthesized and those that already exist but which have not yet been identified) is unimaginably large.

How many distinct organic chemical entities could hypothetically be synthesized and added to a seemingly limitless, ever-expanding chemical universe?

By limiting synthesis strictly to combinations of 30 atoms of just C, N, O, or S, **more than  $10^{60}$  structures are possible !**

Expanding the allowable elements to other heteroatoms (e.g., P and halogens) or deuterium, **the limits to the numbers of possible structures in “chemical space” defies imagination.**

# The Chemical Universe

The largest virtual chemical database yet reported comprises small drug-like molecules - - a total of over 977,000,000 structures (Blum and Reymond, 2009).

Restricted to organic molecules containing fewer than 14 atoms of C, N, O, and S (and limited types of Cl-substituted molecules). Excluded likely substituents such as F, Br, I, P, Si, metals, and most Cl.

Database represents the enormously large numbers of chemicals that could possibly be synthesized just from a very limited spectrum of types of elements and numbers of atoms.

Blum, L. C., and J.-L. Reymond. 2009. 970 Million Druglike Small Molecules for Virtual Screening in the Chemical Universe Database GDB-13. J. Am. Chem. Soc. 131 (25):8732-8733.



# 200 Top-Selling Prescribed Drugs (2006)

1. Lipitor	2. Plavix	3. Zolpidem	4. Humira	5. Synthroid	6. Xarelto	7. Celebrex	8. Lasix	9. Norco	10. Lipitor	11. Synthroid	12. Zolpidem	13. Plavix	14. Lipitor	15. Zolpidem	16. Synthroid	17. Lipitor	18. Plavix	19. Zolpidem	20. Synthroid
21. Tylenol	22. Plavix	23. Zolpidem	24. Humira	25. Synthroid	26. Xarelto	27. Celebrex	28. Lasix	29. Norco	30. Lipitor	31. Synthroid	32. Zolpidem	33. Plavix	34. Lipitor	35. Zolpidem	36. Synthroid	37. Lipitor	38. Plavix	39. Zolpidem	40. Synthroid
41. Tylenol	42. Plavix	43. Zolpidem	44. Humira	45. Synthroid	46. Xarelto	47. Celebrex	48. Lasix	49. Norco	50. Lipitor	51. Synthroid	52. Zolpidem	53. Plavix	54. Lipitor	55. Zolpidem	56. Synthroid	57. Lipitor	58. Plavix	59. Zolpidem	60. Synthroid
61. Tylenol	62. Plavix	63. Zolpidem	64. Humira	65. Synthroid	66. Xarelto	67. Celebrex	68. Lasix	69. Norco	70. Lipitor	71. Synthroid	72. Zolpidem	73. Plavix	74. Lipitor	75. Zolpidem	76. Synthroid	77. Lipitor	78. Plavix	79. Zolpidem	80. Synthroid
81. Tylenol	82. Plavix	83. Zolpidem	84. Humira	85. Synthroid	86. Xarelto	87. Celebrex	88. Lasix	89. Norco	90. Lipitor	91. Synthroid	92. Zolpidem	93. Plavix	94. Lipitor	95. Zolpidem	96. Synthroid	97. Lipitor	98. Plavix	99. Zolpidem	100. Synthroid
101. Tylenol	102. Plavix	103. Zolpidem	104. Humira	105. Synthroid	106. Xarelto	107. Celebrex	108. Lasix	109. Norco	110. Lipitor	111. Synthroid	112. Zolpidem	113. Plavix	114. Lipitor	115. Zolpidem	116. Synthroid	117. Lipitor	118. Plavix	119. Zolpidem	120. Synthroid
121. Tylenol	122. Plavix	123. Zolpidem	124. Humira	125. Synthroid	126. Xarelto	127. Celebrex	128. Lasix	129. Norco	130. Lipitor	131. Synthroid	132. Zolpidem	133. Plavix	134. Lipitor	135. Zolpidem	136. Synthroid	137. Lipitor	138. Plavix	139. Zolpidem	140. Synthroid
141. Tylenol	142. Plavix	143. Zolpidem	144. Humira	145. Synthroid	146. Xarelto	147. Celebrex	148. Lasix	149. Norco	150. Lipitor	151. Synthroid	152. Zolpidem	153. Plavix	154. Lipitor	155. Zolpidem	156. Synthroid	157. Lipitor	158. Plavix	159. Zolpidem	160. Synthroid
161. Tylenol	162. Plavix	163. Zolpidem	164. Humira	165. Synthroid	166. Xarelto	167. Celebrex	168. Lasix	169. Norco	170. Lipitor	171. Synthroid	172. Zolpidem	173. Plavix	174. Lipitor	175. Zolpidem	176. Synthroid	177. Lipitor	178. Plavix	179. Zolpidem	180. Synthroid
181. Tylenol	182. Plavix	183. Zolpidem	184. Humira	185. Synthroid	186. Xarelto	187. Celebrex	188. Lasix	189. Norco	190. Lipitor	191. Synthroid	192. Zolpidem	193. Plavix	194. Lipitor	195. Zolpidem	196. Synthroid	197. Lipitor	198. Plavix	199. Zolpidem	200. Synthroid

# Black Holes in Exploring the Universe Chemical by Chemical

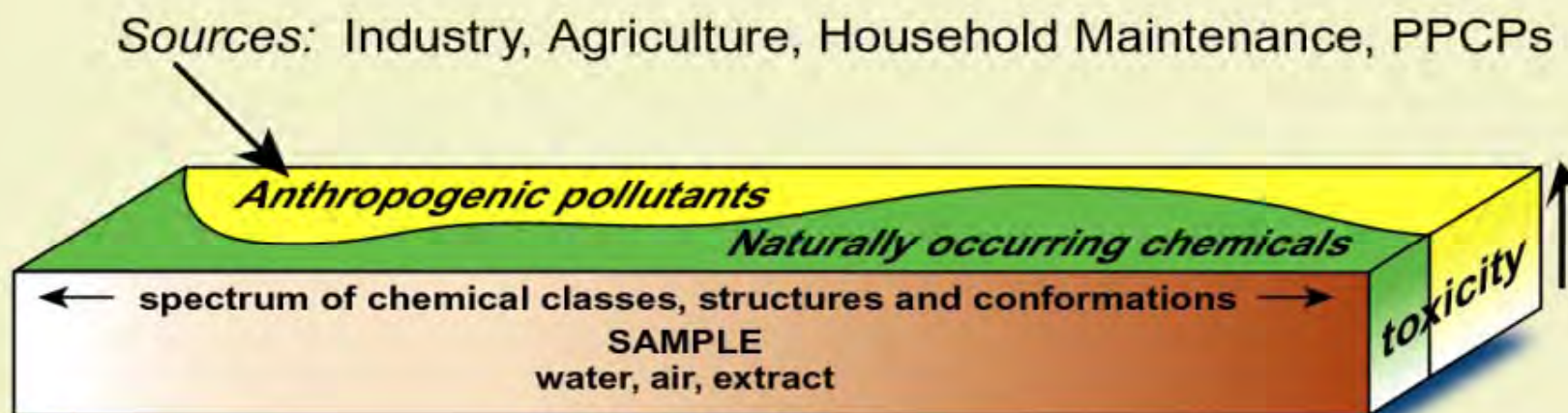


- Traditional chemical-by-chemical approach for evaluating risk of new and existing chemicals is not sustainable (speed, complexity, cost)
- Extensive environmental and toxicological data exist for very few chemicals
- New paradigm will be required to cope with the backlog of existing chemicals and the sheer number of possible new chemicals
- Realization of the critical importance of multi-generational, simultaneous exposure to individual trace levels of multitudes of chemical stressors
- Subtle, delayed-onset effects
- Chemicals cannot be prioritized based solely on commercial volume and conventional toxicity testing
- Assessment of hazard (both virtual and empirical) will require extremely high throughput



**Lessons learned from emerging contaminants will undoubtedly play a major role in the future of risk assessment and risk reduction/mitigation.**

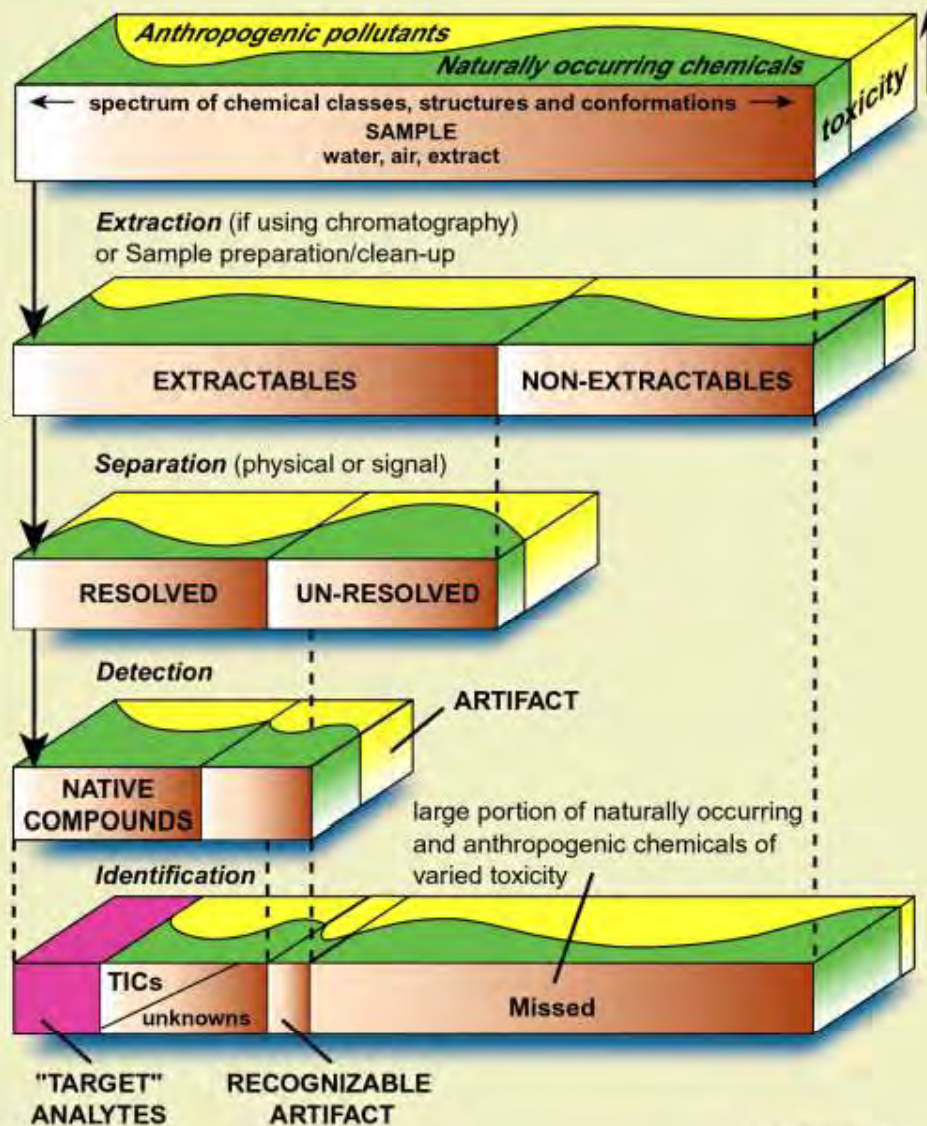
# Universe of Chemicals in the Environment



From: Daughton, C.G. "Cradle-to-Cradle Stewardship of Drugs for Minimizing Their Environmental Disposition while Promoting Human Health. I. Rationale for and Avenues toward a Green Pharmacy," *Environ. Health Perspect.* **2003**, 111(5):757-774; available: <http://www.epa.gov/nerlesd1/bios/daughton/green1.pdf>.



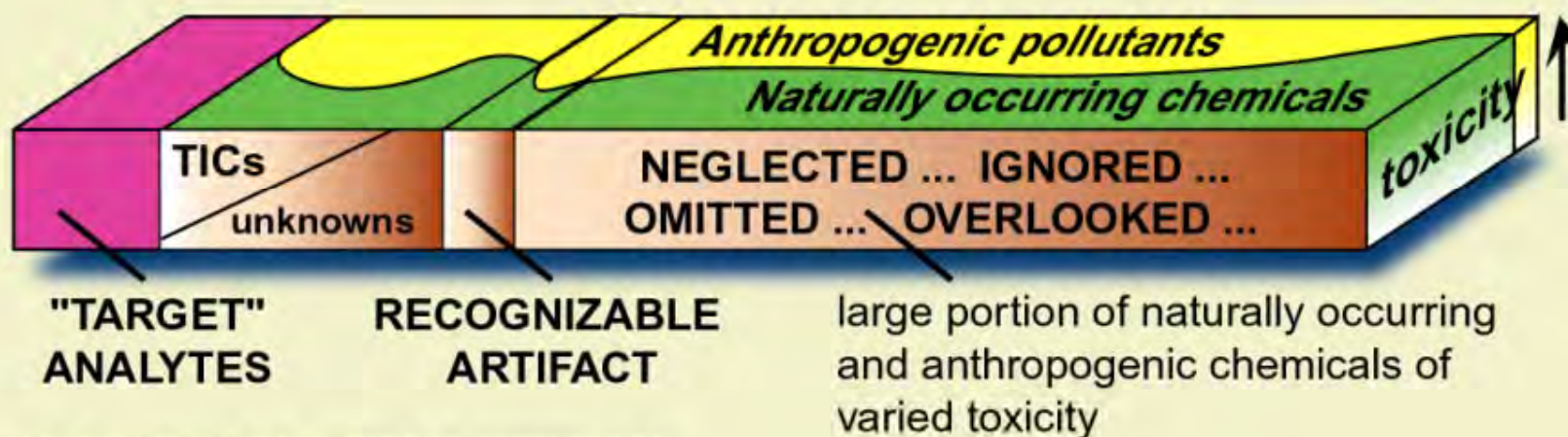
# Limitations and Complexity of Environmental Chemical Analysis



TICs = tentatively identified compounds

C.G. Daughton  
U.S. EPA July 2002

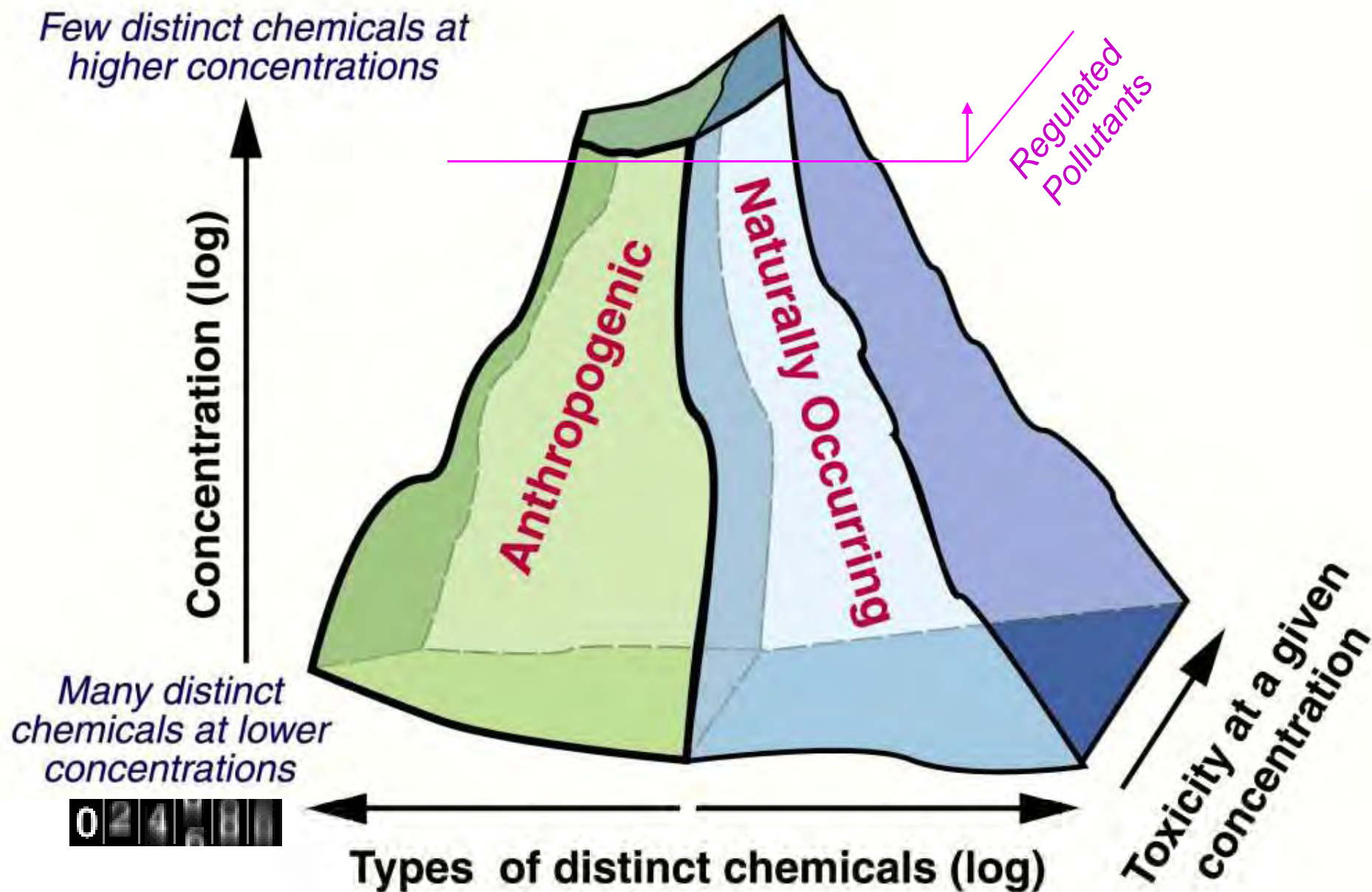
# Chemical Analysis Output for a Typical Environmental Sample



TICs = tentatively identified compounds



# Prevalence/Distribution of Xenobiotic Occurrence

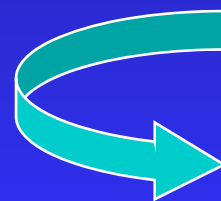


# Einstein on: *Environmental Monitoring*

“Not everything that can be counted counts,  
and not everything that counts can be  
counted.” (oft attributed to Albert Einstein)



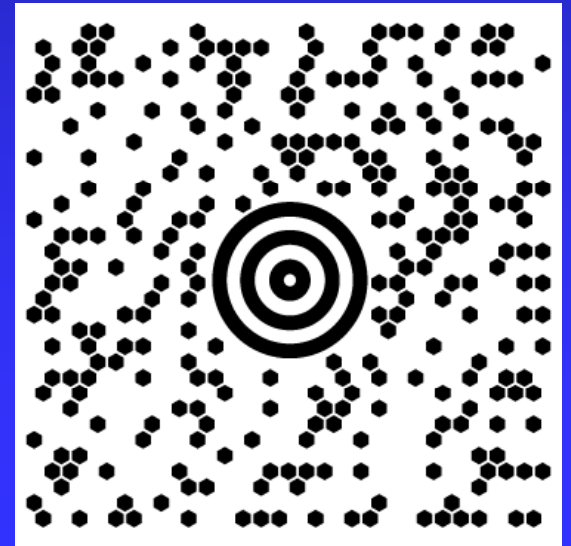
*corollary for environmental monitoring*

 **Not everything that can be measured is  
worth measuring, and not everything  
worth measuring is measurable.**



## further truisms regarding *Environmental Monitoring*

- What one finds usually depends on what one aims to search for.
- Only those compounds targeted for monitoring have the potential for being identified and quantified.
- Those compounds not targeted will elude detection.
- The spectrum of pollutants identified in any sample often represent but a small portion of those present and they are of unknown overall significance with respect to risk.

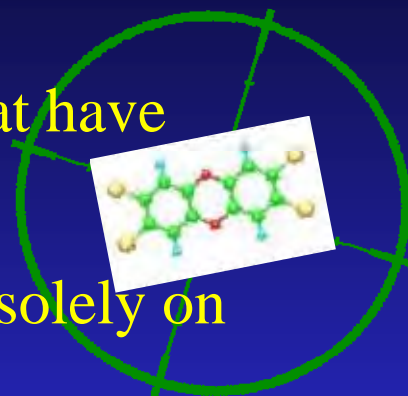




# *Biasing Our Perspective of Chemical Exposure?*

Monitoring studies tend to target those chemicals that have already been reported in prior studies.

Targeted analytes therefore are often self-selected – solely on the basis of prior positive results.

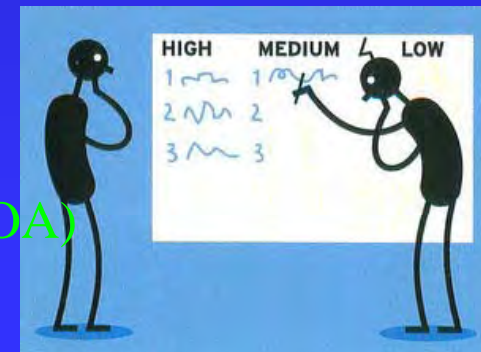


- Are published data providing a highly biased, selective view of the world of chemical contaminants?
- How large is the chemical world that monitoring activities are overlooking?
- How do we establish how accurate and comprehensive the current picture is?

# *Weaknesses in schemes for selecting or prioritizing ECs for monitoring or assessing risk*

**Chemical-by-chemical approaches must account for stressors that:**

- have long been present but are not yet known (chemical unknowns, including natural products and degradates from known chemicals)
- are truly emerging (e.g., new illicit chemicals; by-products from new processes)
- have mechanisms of action not yet recognized - with effects not yet evident (e.g., delayed-onset; subtle, cumulative effects; effects hidden within the variance of natural background)
- have no inherent toxicity of their own but which potentiate the toxicity of others (e.g., efflux pump inhibitors)
- while present at low levels, are extremely potent or can combine for additive or interactive effects (e.g., drugs in same therapeutic class, sharing same MOA)



## *New paradigm needed for addressing ECs?*

Limitations and vulnerabilities of target-based chemical monitoring show that an alternative monitoring approach might be more efficacious:

### **Shift initial focus to biological effects and away from stressors:**

Chemical monitoring places the emphasis on identifying potential stressors to which humans and the environment may or may not ever be exposed - and for which, the potential for effects is largely unknown. *Chemical monitoring often appears to become the end in itself - creating a vacuum for communicating what it might mean.* The historic focus has been on the stressor rather than on outcome or ultimate concern - namely, the potential for exposure and biological effects.

### **Biological endpoints to inform and guide monitoring:**

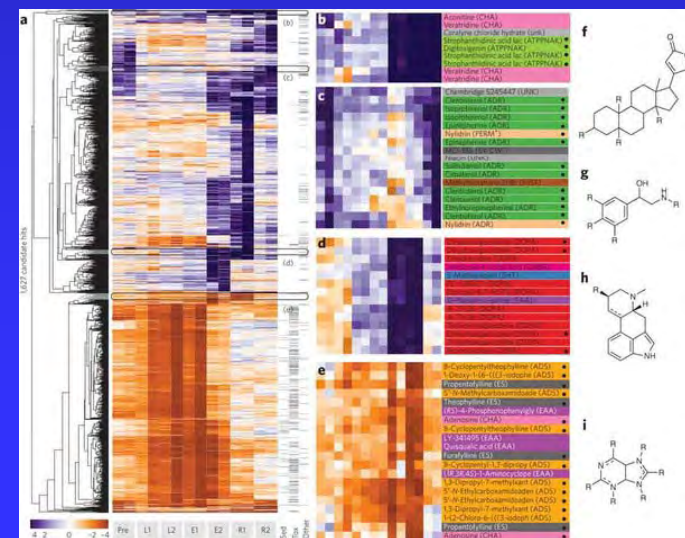
Given the sheer numbers of ECs that might exist now and in the future, might the power and meaning of target-based chemical monitoring be vastly improved by instead using biology-based screening as the initial survey tool to guide subsequent chemical monitoring?



# New paradigm needed for addressing ECs?

## Example:

Holistically assess overall potential for adverse biological impact from chemical contaminants with standardized arrays of biology/biochemical assays based on evolutionarily conserved cellular processes whose responses integrate the signals from all chemical stressors present. Examples include cell- or biochemical-based assays (or biomarkers) for measuring inhibition of efflux pumps, endocrine modulation responses, and cellular stress responses.



## *New paradigm needed for addressing ECs?*

Might protecting human health and the environment be served more directly and efficiently by:

Empowering the public: Complexity of the EC issue poses extreme challenges for communication of risk to the public - and prevention or misperception of risk. Critical to involve other disciplines (particularly social psychologists) in the ongoing dialog regarding chemical pollution. Better public understanding of chemical pollution could dramatically facilitate its reduction or elimination.

Designing a sustainable chemocentric society: More efficient use of chemicals and wider implementation of green chemistry practices integrated with pollution prevention and stewardship.

Regardless of whether ECs are ever shown to pose hazards as trace environmental pollutants, actions directed at reducing their entry to the environment via stewardship and pollution prevention (coupled with heightened public awareness) could achieve immediate outcomes. Leveraging would inevitably result, yielding collateral beneficial outcomes extending far beyond those targeted specifically at ECs.



# *Are we headed in the right direction? What should we expect to achieve?*



- Are our priorities in balance with society's expectations and needs?
- Are we focusing on those areas that will yield the most important outcomes?
- Is research sufficiently coordinated and leveraged - internationally?



## *Possible Major Outcomes from Emerging Contaminants Research*

- Improve our ability to assess risk more comprehensively.
- Expand the known universe of chemical stressors. Provide means to identify previously unrecognized or unexpected contaminants.
- Improve the public's understanding of risk - and how all actions can impact the environment.



## *Possible Major Outcomes from Emerging Contaminants Research*

- Catalyze new paradigm for screening and controlling chemical unknowns in complex mixtures that pose actual rather than potential risks.
- Reduce cost and improve speed and efficiency of managing, mitigating, and preventing chemical pollution.
- Develop proactive capability for early warning or predictive ability for identifying newly emerging contaminants.



# Questions to Consider



- Is it useful to call well-established ubiquitous environmental contaminants "emerging"?
- Would a term other than ECs be more accurate and useful for "unregulated" chemical contaminants?
- Should such a term also convey a clear meaning for the public and regulators?
- How might progress and success be assessed with regard to research in ECs?



## *Questions to Consider*

- Can a process be designed to ensure that today's "emerging contaminants" don't persist, becoming tomorrow's concern?
- At what point can a chemical be "graduated" out of the domain of "emerging"?
- Can a proactive system be designed that obviates the need for the term "emerging" in the first place?
- Can stewardship and green chemistry programs be designed to effectively prevent the entry of ECs to the environment?



## *Final Thoughts*



- An individual chemical contaminant can be viewed in a context beyond that of its specific molecular structure and particular toxicological properties.
- Ideally, its presence in the ambient environment might be expected (at the worst) to comport with its value in sustainable use.
- Focus has long been on the individual chemical as an environmental contaminant rather than on preventing its becoming a contaminant in the first place. A sustainable balance might exist between the manufacture and the ultimate uses of any given chemical.

# *Final Thoughts*



- The types and trends in levels of chemical contaminants in the environment serve as direct measures of our progress toward sustainability - from green and virtual chemistry, to smart manufacturing and intelligent consumption.
- Sustainability could be advanced from consumers better understanding their individual roles in both causing (via combined miniscule inputs) and preventing the release of chemicals to the environment. ECs serve to highlight that the environment and humans are intricately interconnected in a complex system that would benefit from examination as an integral whole.



*feel free to contact:*

**Christian Daughton, Ph.D.**

Chief, Environmental Chemistry Branch  
Environmental Sciences Division  
National Exposure Research Laboratory  
U.S. Environmental Protection Agency  
Las Vegas, Nevada 89119

*daughton.christian@epa.gov*

*702-798-2207*

<http://www.epa.gov/nerlesd1/bios/daughton.htm>



*prepared for:*

***NorCal SETAC 20<sup>th</sup> annual meeting***  
**Contaminants of Emerging Concern:**  
**Responding to a 21st Century Challenge**

University of California, Berkeley

13 May 2010

**Christian Daughton, Ph.D.**

Chief, Environmental Chemistry Branch  
Environmental Sciences Division  
National Exposure Research Laboratory  
U.S. Environmental Protection Agency  
Las Vegas, NV 89119  
daughton.christian@epa.gov  
702-798-2207

prepared: 3 May 2010

Office of Research and Development  
National Exposure Research Laboratory, Environmental Sciences Division, Las Vegas, Nevada